Reply to Office Action of June 17, 2004 Amendment No. 6 dated August 24, 2004

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application:

Listing of Claims:

- 1. (Previously presented) Method for the preparation of an antibodytumor cell preparation for immunization of humans and animals against tumor cells comprising the steps of:
 - a) isolating autologous tumor cells;
- b) treating the tumor cells to prevent the survival thereof following reinfusion;
- c) incubating the thus treated tumor cells with intact heterologous bispecific antibodies showing the following properties:
 - (i) binding to a T cell;
 - (ii) binding to at least one tumor-associated antigen on a tumor cell;
 - (iii) binding, by their Fc portion to Fc receptor-positive cells; and
 - (iv) capable of activating the Fc receptor-positive cell whereby the expression of cytokines, co-stimulatory antigens or both is induced or increased,

wherein the bispecific antibodies have isotype combinations selected from the group consisting of:

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rat-IgG2b/human-IgG1,
rat-IgG2b/human-IgG3,
rat-IgG2b/human-IgG3[oriental allotype G3m(st) = binding to protein A],
rat-IgG2b/human-IgG4,
rat-IgG2b/rat-IgG2c,
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mouse-IgG2a/human-IgG3[caucasian allotypes G3m(b+g) = no binding to protein A, in the following indicated as *],

mouse-IgG2a/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

mouse-IgG2a/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human- IgG3*-[CH2-CH3],

mouse-IgG2a/human-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

mouse-[VH-CH1,VL-CL]-human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

mouse-[VH-CH1,VL-CL]-human-IgG4/rat-[VH-CH1,VL-CL]-human-IgG4-[hinge]-human-IgG4[N-terminal region of CH2]-human-IgG3*[C-terminal region of CH2: > aa position 251]-human-IgG3*[CH3],

rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge-CH2-CH3],
rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG2-[hinge-CH2-CH3],
rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG3-[hinge-CH2-CH3,
oriental allotype],

rat-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG4-[hinge-CH2-CH3], human-IgG1/human-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG4[N-terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position 251]-human-IgG3*[CH3],

human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG4[N-terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position 251]-human-IgG3*[CH3],

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human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG2[N-terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position 251]-human-IgG3*[CH3],

human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG2[N-terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position 251]-human-IgG3*[CH3],

human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

human-IgG2/human-[VH-CH1,VL-CL]-human-IgG2-[hinge]-human-IgG3*-[CH2-CH3],

human-IgG4/human-[VH-CH1,VL-CL]-human-IgG4-[hinge]-human-IgG3*-[CH2-CH3],

human-IgG4/human-[VH-CH1,VL-CL]-human-IgG4-[hinge]-human-IgG4[N-terminal region of CH2]-human-IgG3*[C-terminal region of CH2 : > aa position 251]-human-IgG3*[CH3],

mouse-IgG2b/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

mouse-IgG2b/human-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

mouse-IgG2b/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG3*-[CH2-CH3],

mouse-[VH-CH1,VL-CL]-human-IgG4/rat-[VH-CH1,VL-CL]-human-IgG4-[hinge]-human-IgG4-[CH2]-human-IgG3*-[CH3],

human-IgG1/rat-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG4-[CH2]-human-IgG3*-[CH3],

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human-IgG1/mouse-[VH-CH1,VL-CL]-human-IgG1-[hinge]-human-IgG4-[CH2]-human-IgG3*-[CH3],

human-IgG4/human-[VH-CH1,VL-CL]-human-IgG4-[hinge]-human-IgG4-[CH2]-human-IgG3*-[CH3],

rat-IgG2b/mouse-IgG2a, rat-IgG2b/mouse-IgG2b, and rat-IgG2b/mouse-IgG3.

- 2. (Previously presented) Method according to claim 1, in which said antibodies are selected so that they are capable of binding Fc receptor-positive cells having a Fcy receptor I, II, or III.
- 3. (Previously presented) Method according to claim 2, in which said $Fc\gamma$ receptor I-positive cells are selected from the group consisting of monocytes, macrophages, dendritic cells, and activated neutrophils.
- 4. (Previously presented) Method according to claim 1, in which said antibodies are capable of inducing tumor-reactive complement-binding antibodies and thus inducing a humoral immune response.
- 5. (Previously presented) Method according to claim 1, in which said antibodies are selected to bind to the T cells via CD2, CD3, CD4, CD5, CD6, CD8, CD28 or CD44.
- 6. (Previously presented) Method according to claim 1, in which said antibodies are selected so that following their binding to the Fc receptor-positive cells the expression of CD40, CD80, CD86, ICAM-1 and/or LFA-3 as co-stimulatory antigens, and/or secretion of cytokines by the Fc receptor-positive cell is initiated or increased.

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- 7. (Previously presented) Method according to claim 1, in which said antibodies are selected so that the secretion of IL-1, IL-2, IL-4, IL-6, IL-8, IL-12 being cytokines or of TNF-α or a combination thereof is increased.
- 8. (Previously presented) Method according to claim 1, in which said bispecific antibody is selected to be an anti-CD3 X anti-tumor-associated antigen antibody or anti-CD4 X anti-tumor-associated antigen antibody or anti-CD5 X anti-tumor-associated antigen antibody or anti-CD8 X anti-tumor-associated antigen antibody or anti-CD2 X anti-tumor-associated antigen antibody or anti-CD2 X anti-tumor-associated antigen antibody or anti-CD44 X anti-tumor-associated antigen antibody.

9-12. (Canceled)

- 13. (Previously presented) A method for preparing a vaccine comprising an antibody-tumor cell preparation, said method comprising preparing an antibody-tumor cell preparation by the method of claim 1, and preparing a vaccine from said antibody-tumor cell preparation.
- 14. (Previously presented) A method for preparing a vaccine comprising activated peripheral blood mononucleated cells, said method comprising preparing an antibody-tumor cell preparation by the method of claim 1 in which step (c) is replaced with step (d), which comprises incubating the thus-treated tumor cells with both said intact heterologous bispecific antibodies and peripheral blood mononucleated cells, thereby activating said peripheral blood mononucleated cells, and preparing a vaccine from the thus-activated peripheral blood mononucleated cells.
- 15. (Previously presented) Method according to claim 1, in which said tumor cells are incubated with the antibodies for a period of 10 minutes to 5 hours.

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- 16. (Previously presented) Method according to claim 1, in which said tumor cells are incubated with the antibodies for a period of 15 minutes to 120 minutes.
- 17. (Currently amended) Method according to claim 14 in which said incubation of step (e) (d) is performed for a period of 1 to 14 days.
- 18. (Currently amended) Method according to claim 14 in which said incubation of step (e) (d) is performed with about 10⁸ to 10¹⁰ mononucleated peripheral cells.
- 19. (Previously presented) Method according to claim 1, in which said tumor cells are present in the amount of about 10⁷ to 10⁹ cells.
- 20. (Previously presented) Method according to claim 1, in which said bispecific antibodies are added in an amount of 2 to 100 μg.
- 21. (Previously presented) Method according to claim 1, in which said treating of the tumor cells in step b is performed by irradiation.
 - 22. (Canceled)
- 23. (Currently amended) Method for preventing the reoccurrence of a tumor eaused by the tumor cells against which the intact heterologous bispecific antibodies of claim 1 are directed, said method comprising administering to an individual in whom such tumor cells have appeared a an antibody-tumor cell preparation prepared according to the method of claim 1.

24-25. (Canceled)

26. (Currently amended) A pharmaceutical composition comprising a an antibody-tumor cell preparation obtained by the method of claim 1.

27-30. (Canceled)

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- 31. (New) A pharmaceutical composition comprising the activated peripheral blood mononucleated cells obtained by the method of claim 14.
- 32. (New) A method for preventing the recurrence of a tumor, said method comprising: preparing an antibody-tumor cell preparation by the method of claim 1 in which step (c) is replaced with step (d), which comprises incubating the thus-treated tumor cells with both said intact heterologous bispecific antibodies and peripheral blood mononucleated cells, thereby activating said peripheral blood mononucleated cells; and administering to an individual in whom such tumor cells have appeared the activated peripheral blood mononucleated cells.
- 33. (New) The method of claim 32, wherein the peripheral blood mononucleated cells are added following a preincubation of the thus-treated tumor cells with said intact heterologous bispecific antibodies.
- 34. (New) The method of claim 14, wherein the peripheral blood mononucleated cells are added following a preincubation of the thus-treated tumor cells with said intact heterologous bispecific antibodies.
- 35. (New) A pharmaceutical composition comprising the activated peripheral blood mononucleated cells obtained by the method of claim 32, 33, or 34.